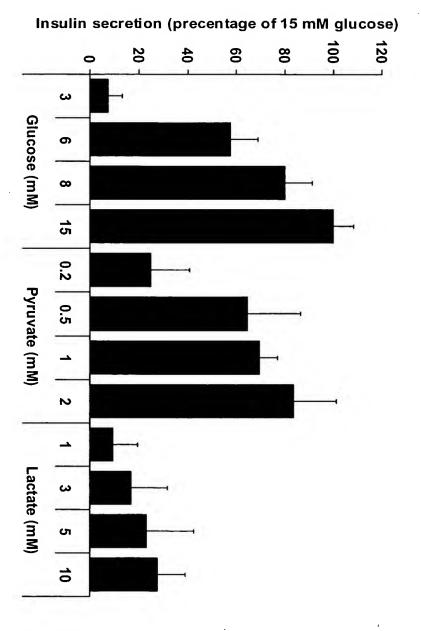


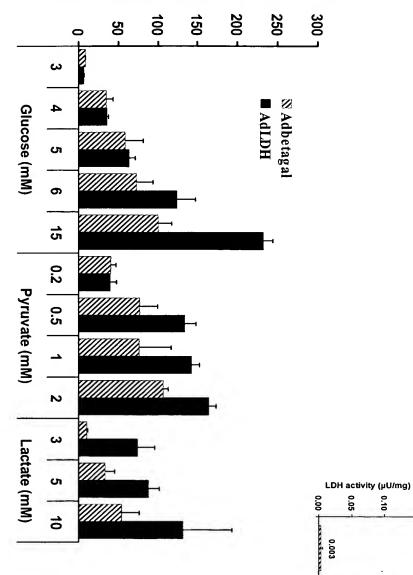
Fig. 1

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SSVPVWSGVNVAGVSLKSLNPQLGTDADKEQWKDVHKQVVDSAYEVIKLKGYTSWAIGLSVADLA QHGSLFLKTPKIVSSKDYSVTANSKLVIITAGARQQEGESRLNLVQRNVNIFKFIIPNVVKYSPQ MAALKDQLIVNLLKEEQVPQNKITVVGVGAVGMACAISILMKDLADELALVDVIEDKLKGEMMDL ESIMKNLRRVHPISTMIKGLYGIKEDVFLSVPCILGQNGISDVVKVTLTPDEEARLKKSADTLWG CKLLIVSNPVDILTYVAWKISGFPKNRVIGSGCNLDSARFRYLMGERLGVHPLSCHGWVLGEHGD IQKELQF



Insulin secretion (as presentage of 15mM)





0.093

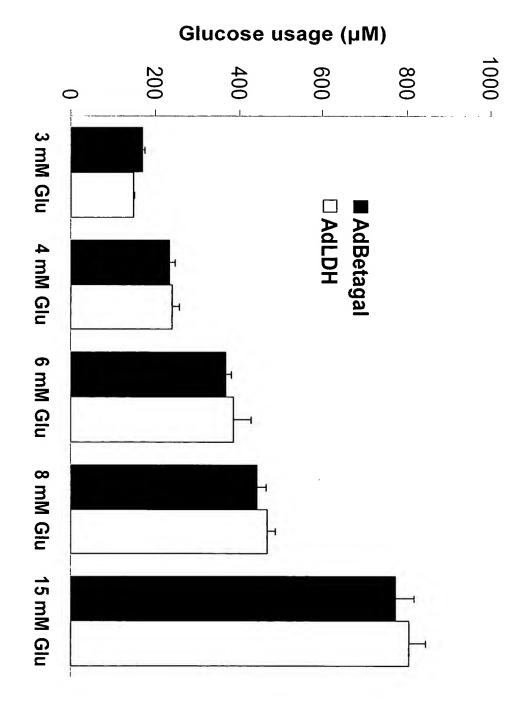
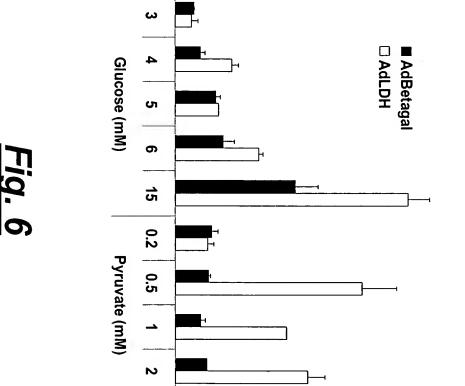


Fig. 5



100 -

Lactate output

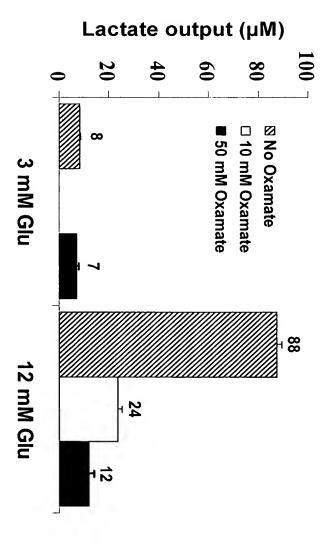
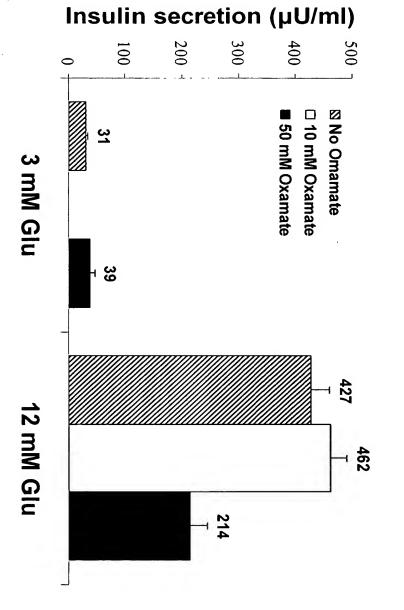


Fig. 7A

Insulin secretion



ig. 7B

Pyruvate Cycling

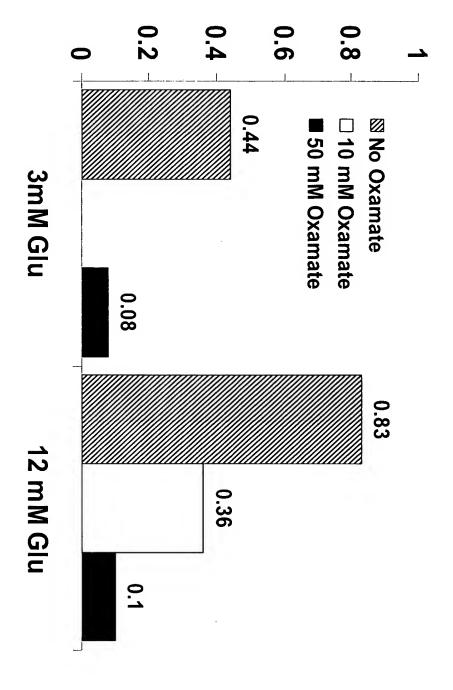


Fig. 7C

Oxamate inhibits Insulin secretion in islets

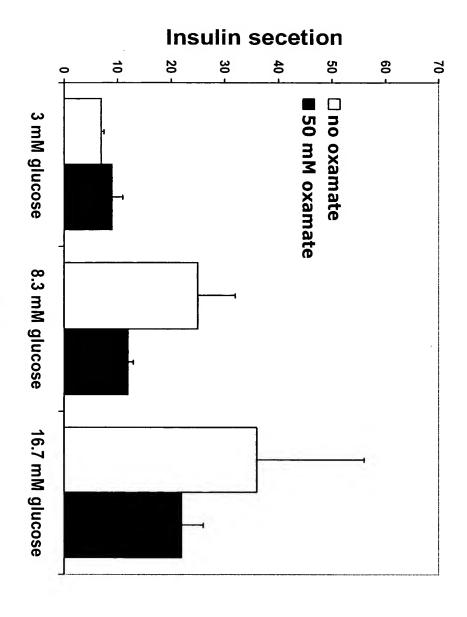
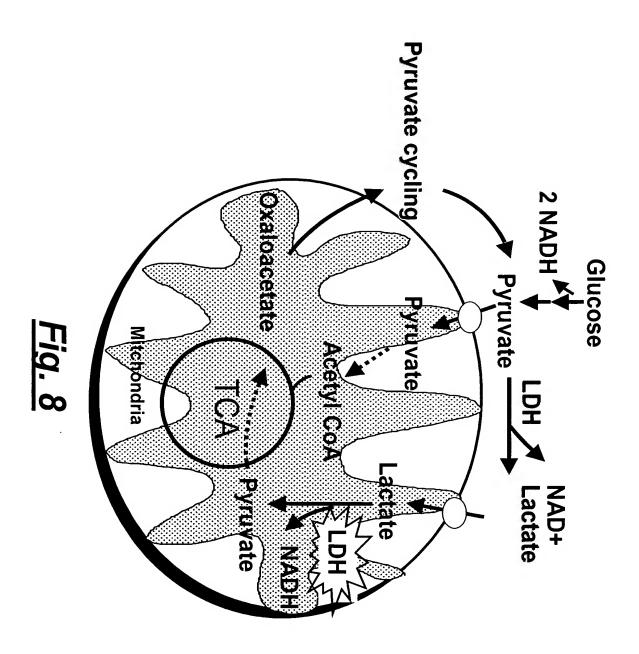
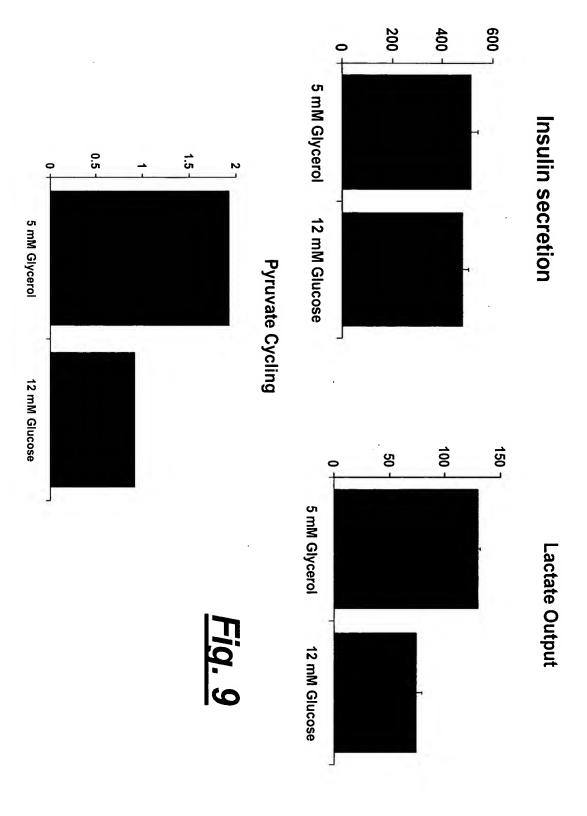


Fig. 7D



Production in Glycerol Kinase Expressing Cells. Pyruvate Cycling Correlates with Lactate



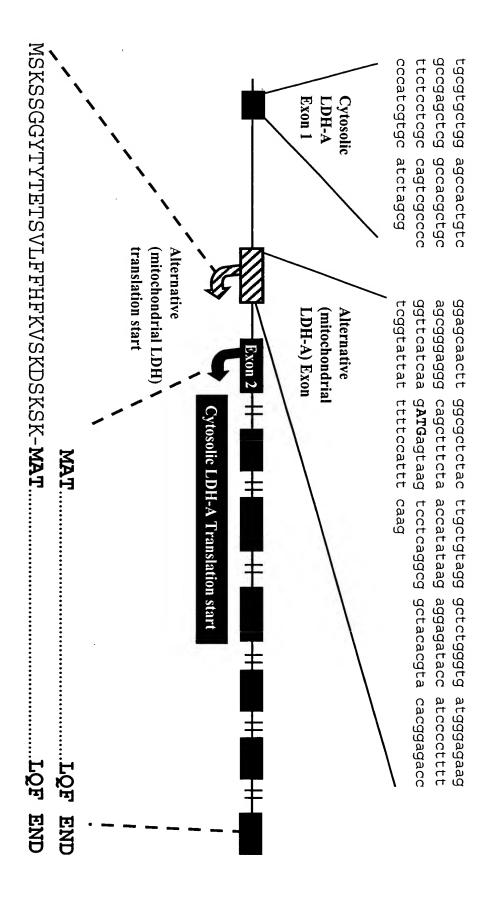


Fig. 10

among different species The N-terminal leader sequence is conserved

- Rat **MSKNSGGYTYTETSVLFFHFKVPKDSKSK**
- Mouse MSKSSGGYTYTETSVLFFHFKVSKDSKSK
- Human MGEPSGGYTYTQTSIFLFHAKIPFGSKSN
- MSK SGGYTYTETSVLFFHFKVPKDSKSK

CCCTTTTGGG GTTCATCAAG ATGAGTAAGA ACTCAGGCGG CTACACATAT ACGGAGACCT CAGTATTATT TITCCATTTC

AAGGTCCCAA AAGATTCAAA GTCCAAG**ATG** GCAGCCCTCA AGGACCAGCT GATTGTGAAT CTTCTTAAGG AAGAACAGGT

GAGGAGGCCC AAGCAGGTGG GGGAGTTCAT GGCTTCCCCA CCGTCAGCAA CCTAGCACTT AGGATGTCTT CTTGGCCGAG ACGTCGCCGG TGAAATACAG CTTAAGACAC CTGATGAGCT CCCCCAGAAC AAGATTACAG CACTGTCCAG GCTGCAGCAG GGTTTCTATG GAGACCACGC ACTTCTCATC TGAGCTGTGG TTAGTCCAGT GCCTGAAGAA CCTCAGCGTC AGCATAATGA CCACTGAGCT AAAACAAAGT GAGGGAGAGA TTGACAGTGC CAAAAATTGT TGCCCTTGTT CGTCTCCCTG TCCACAGTGC GAGTGCAGAT CCATGTATCC AGAACCTTAG ATACGAAGTG GTCACGGGTG TATTGGAAGT GATGTCATAG TIGTIGGGGT AAGTCTCTGA AAACTGCTCA GCCGGCTCAA CTCCAGCAAA ACCCTCTGGG TGGGACAAAA GCGGGTGCAT ATCAAGCTGA GGTCCTGGGA GGTTGCAATC TCGTCTCAAA GATTATAGTG AAGATAAGCT TGGTGCTGTT ACCCGCAGCT TTTGGTCCAG GAATCCAGAA TGGAATCTCA GAGCATGGCG CCCATTTCCA AAGGTTACAC GGGCACGGAT TGGATTCGGC CCCAGTGGAT CGAAACGTGA AAAGGGAGAG GGCATGGCTT TGACTGCAAA GGAGCTGCA G GTGCCATCA G GATGTTGTG A CCATGATTA A ATCCTGGGC C GCAGACAAG G ACTCCAGTG T TCGGTTCCG T ATCTTGACC T ACATCTTCA A CTCCAAGCT G ATGATGGAT C GGGTCTCTAT TACCTGATGG TTCTAAAGTC AGGTGACACT GCCTGTGTGG GTTCATCATT GTCATTATCA ATTGGCCTCT AGCAGTGGAA ACGIGGCTIG TTCAGCATGG TATCTTAATG GACTCCTGAC GGGATCAAGG CCGTGGCAGA AGTGGTGTGA GAGAAAGGCT CCAAATGTTG CAGCCTTTTC AAGGACTTGG GGATGTGCAC GAAGATCAGC CCGCGGGGGC TICCCAGIGI

- Mitochondrial start site
- \$ cytosolic start site
- Overlined 5'ORF
- Primer sequences are underlined

MSKNSGGYTYTETSVLFFHFKVPKDSKSKMAALKDQLIVNLLKEEQVPQNKITVVGVGAVGMACA SARFRYLMGERLGVHPLSCHGWVLGEHGDSSVPVWSGVNVAGVSLKSLNPQLGTDADKEQWKDVH EGESRLNLVQRNVNIFKFIIPNVVKYSPQCKLLIVSNPVDILTYVAWKISGFPKNRVIGSGCNLD QNGISDVVKVTLTPDEEARLKKSADTLWGIQKELQF KQVVDSAYEVIKLKGYTSWAIGLSVADLAESIMKNLRRVHPISTMIKGLYGIKEDVFLSVPCILG ISILMKDLADELALVDVIEDKLKGEMMDLQHGSLFLKTPKIVSSKDYSVTANSKLVIITAGARQQ

ATATATTATATGTGTCTGTAGTGTGCATTGCAATATTATGTGAGATGTAAGATCTGCATATGGA TGATGGAACCAACCACCCAAGTG CCAGTGTGCCTGTGTGGAGTGGTGTGAATGTTGCCGGCGTCTCCCTGAAGTCTCTTAACCCAGA ACTGGGCACTGACGCAGACAAGG ATCCAGTGGATATCTTGACCTACGTGGCTTGGAAAATCAGTGGCTTTCCCAAAAACCGAGTAAT TGGAAGTGGTTGCAATCTGGATT TGGTCCAGCGAAACGTGAACATCTTCAAGTTCATCATTCCCAACATTGTCAAGTACAGTCCACA CTGCAAGCTGCTGATCGTCTCCA GCAAAGACTACTGTGTAACTGCGAACTCCAAGCTGGTCATTATCACCGCGGGGGCCCCGTCAGCA AGAGGGGGAGAGCCGGCTCAACC TCATGCCAAATAAAACCTTGAACAGTG CGAGTTCCCAGTTAAGTCGTATAACCTGGCTCCAGTGTGTACGTCCATGATGCATATCTTGTGC ATAAATGTTGTACAGGATATTTT ${ t GTCTCTGAGACACTGCCAACTGCAGGCTTCGATTACCCCTGTGAGCCTGCTGCATTGCTGCC$ TGGGGAAACATCTCACTCCCCACAGCTCTGCCCTGCTGCCAAGTGGTACTTGTGTAGTGGTGAC CTGGTTAGTGTGACAGTCCCACT TTCACTGTCCAGGCTGCAGCAGGGCTTCTAGGCAGACCACACCCTTCTCGTCTGAGCTGTGGTT AGTACAGTGGTGTTGAGATGGTG AAGAGGCCCGCCTGAAGAAGAGCGCAGACACCCTCTGGGGAATCCAGAAGGAGCTGCAGTTCTA AAGTCTTCCCCGTGTCCTAGCAC TCTCTGTGGCAGACTTGGCTGAGAGCATAATGAAGAACCTTAGGCGGGTGCATCCCATTTCCAC CATGATTAAGGGTCTCTATGGAA AGCAGTGGAAGGAGGTTCACAAGCAGGTGGTGGACAGTGCCTACGAGGTGATCAAGCTGAAAGG TTACACATCCTGGGCCATTGGC CAGCGCGGTTCCGTTACCTGATGGGAGAGAGGCTGGGGGTTCACGCGCTGAGCTGTCACGGCTG GGTCCTGGGAGAACATGGCGACT AGATTACAGTTGTTGGGGGTTGGTGCTGTTGGCATGGCTTGTGCCATCAGTATCTTAATGAAGGA CTTGGCGGATGAGCTTGCCCTTG CCATCCCCTTTTGGTTCATCAAGATGAGTAAGTCCTCAGGCGGCTACACGTACACGGAGACCTCGGTATTATTTTTCCATTTCAAGG ${ t TCAATGAGGATGTCTTCCTCAGTGTCCCCATGTATCCTGGGACAAAATGGAATCTCGGATGTTGTGAAGGTGACACTGACTCCTGAGG$ TTGACGTCATGGAAGACAAACTCAAGGGCGAGATGATGGATCTCCAGCATGGCAGCCTCTTCCT TAAAACACCAAAAATTGTCTCCA ${ t TCTCAAAAGATTCAAAGTCCAAGATGGCAACCCTCAAGGACCAGCTGATTGTGAATCTTCTTAA GGAAGAGCAGGCTCCCCAGAACA$

SARFRYLMGERLGVHALSCHGWVLGEHGDSSVPVWSGVNVAGVSLKSLNPELGTDADKEQWKEVH MSKSSGGYTYTETSVLFFHFKVSKDSKSKMATLKDQLIVNLLKEEQAPQNKITVVGVGAVGMACA QNGISDVVKVTLTPEEEARLKKSADTLWGIQKELQF KQVVDSAYEVIKLKGYTSWAIGLSVADLAESIMKNLRRVHPISTMIKGLYGINEDVFLSVPCILG EGESRLNLVQRNVNIFKFIIPNIVKYSPHCKLLIVSNPVDILTYVAWKISGFPKNRVIGSGCNLD ISILMKDLADELALVDVMEDKLKGEMMDLQHGSLFLKTPKIVSSKDYCVTANSKLVIITAGARQQ

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136 136 156	DADATOBOOBASTDAABAATBABTABAASTASTTSS BTTT-TSSST DATATSSABASSASTASTASTTSS BTTT-TSSST DATATSSABASSASTASTASTASTASTASTASTASTASTASTASTASTA	٧8	ratmitlDHcloned humanmitlDHmRNA mousemitlDHmRNA	
06 98 64	ADDABABABATUTDDDAATTTDTTBADBBBBBDDBABAA - BAB ADDBBAAABBATCTTTTTATADABADATD ABBDDBABATDTD ADDATABBBBABAATATADDAAT DTTTDBADBBBBDBBABAA - BAB *** *** * * * * * * * * * * * * * * *	39	ratmitLDHcloned humanmitLDHmRNA mousemitLDHmRNA	
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Alignment Mode: Slow Alignment Parameters: Gap Penalty = 10.0 Extend Gap Penalty = 5.0			Pairwise Alignme	
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832	SABAADTOTOTOTOTOTOTTOTAADTAADTAADTATOTOTOTO	987.	ANAmHOLJimasmuh
825	STBAABTOOSTSTBOBBOOBSTBOAABTBSTBABBTBTBTBTBTBTBTBT		ratmitLDHcloned
		722	F-4-[-110 14;-4
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S84	COLOROGO DE LA COMPANA DE LA CO	984	моизетісърнивия
58L	TTCACCCATTAAGCTGTCATGGGTGCTCCTTGGGGAACATGGAGATTCC	987	АИЯшНСЛЭ і шивти
SLL	COTOADOBTACABABBBTCOTBBBTCACTTTTACTT		ratmitLDHcloned
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5£7	DADDDTDDDAAADDDDDTADTDDATTDDDTTADDDDADTTADDTDTAAD	989	AMAmHddtimamud
725	DADOTTOGGTTGGTTATTCCTTATTCGGTTGGAAAAGCTGGGAG	949	ratmitLDHcloned

989	STT99T9AA99TTAAT9A9>>AAAAA>>>TTT99AGAAAAA99TT9	959	тои зе ті СПР Н т КИ Р
589	GCTTGGAAGATAATTTCCCAAAAACCGTGTTTTGGAAGATGGTTG		humanmitLDHmRNA
S L 9	GCTTGGAAGATTATTGGAAAAAACCACATTGGAAAGTTGGTTG		ratmitLDHcloned
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932	PT9DATDDADTTDTATA99T9ADDTAADDTDTA9TD9AAD9TDA	985	АИЯшНСІЗітэгиот
932	DTDDATTDTATADDTDADDTAADTTTDTTATTDTTTDAADDTDA	985	АИЯшНСТЭ і тавтин
929	PTPOATOOAPTTOTATABBTPAOOOAAAOTOTBOTAOTOBTOAAAOBTBA	945	ratmitLDHcloned
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585	CGTGAACATATATATATATATATATATATATATATATATA		ANAmHGLJimnsmud
SLS	DADDTDADATAAADTTTTAAADDTTADTADTTDAADTTDAAADTDD		ratmitLDHcloned
		761	Pono[DIII] 14; m4ox
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533	AATĐOĐAOOTĐĐTTTAATTOTĐOĐĐAAAĐAĐĐĐAAACĐAĐTĐOĐĐĐĐ	98₺	humanmitlDHmRNA
225	AAADDDADTTTAADTDDDDDDADADADADADADADADDDDDD	9∠₹	ratmitLDHcloned
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121		361	podo[pHd]+;mtex
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5Eħ	5TOTOTETAGAAAOOACAATTOOTTTTOOGAOOSTAGAACOTOTAGO	386	humanmitLDHmRNA
45 E	ADDIDITAAAAADDADAAATTDTTTTDDADDADADTTDTABD	975	ratmitLDHcloned
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385	TADTADADADADADADADADATCTCATCATCATCATCATCATCATCATCATCATCATCAT		humanmitlDHmRNA
	TAUTAUAUAUUAAA TUUAATAUAAAATAUTUTAUTTUTTUUUTTUAA		•

Fig. 13B

Fig. 13C

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BIEI	TOTAD-AAADDDDTDTDDTADADTTDTD-D-TDADATDATTDDTDTD	97.ZT	шолаештсгрншкия
	TOAAOTADAAAAAABOCTGGGAAAAATTTTATATAATDADAAATTATATATATATAT		•
1290	DAAADDTB-BTTBADDTDTD	5961	bano[pHG.11 imtex

727 <i>4</i>	GAGTOTDDT-DTT-DDDADADADADBATOTTDBDADBAD	1532	MousemitLDHmRNA
1281	CAACAGGATT-CTAGGTGG-AGGTTGTGCATGTTGTTGTTGTTGTTGTTGTTGTTGTTGTTGTTGTTGT	1534	humanmitlDHmRNA
T 564	CAGCAGGGTTTCTATGG-AGACCACGCAC-TTC-TCATCTGAG	1552	ratmitLDHcloned
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₽82T	CTGCAGTTCTAAAGTCTTCCCCGTGTCAACTCTCACTGTCACTGTCACGCTG	CRTT	моизептстришкия
			humanmitLDHmRNA
			· ·
VCCL	OTGCAGTTCAACTTCACGAGTCTTCACCTTCACTTCACGTTC	2711	bano[pHC.11 imtex
	****** ***** ***** ** ** ** ** ****** ****		
	PGGCCTGAAGAAGAGAGAGAGAGAAGAAGAAGAAGAGAGAG		<u>-</u>
	DADDAAAACCTATGGGGATACACATACACAAAAAAAAAAA		
₽LTT	DADDAADADTOTOTOTOTOTOTOTOTOTOTOTOTOTOTOT	IISS	ratmitLDHcloned
	* ** *** **** ***** ******** ** ** *****		
TT3	DAADDADTCTCAGTGAAGTGTGAAGTGTGAAGTTGAAGTGAAG	580T	MOUSEMITLDHMRNA
713 4	ACAGAATCTCAGACCTTGTGAAGGTGACTCTGAGGAAG	580T	литапті t L D Н т К И А
	DOADCACTCACACTGTGAAGTGACACTGACTCACAGG		
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TOSŧ	T999AATTA9TACCACCATTT-ACCCTACGEGGGGGGTACACGAGAGAAGTAA	946	ratmitLDHcloned
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S86	${\tt TATAGAGGCATTTGCACTTTTTGCACTTTTTGCACTTTTTGCACTTTTTGCACTTTTTGCACTTTTTGCACTTTTTTTT$	986	humanmitLDHmRNA
S L 6	TACAAGCCGAGCCTCTCCTCTCAGACGCAGCATCTCACACAT	976	ratmitLDHcloned
	**** ** ****** ** ** ** ** ** ** ** **		
586	TTCACAAGTGGAAGTGCCTACGAGGTGATCAAGGTTGAAGGTTGAAGGTTT	988	mousemitLDHmRNA
932	DDDAAACTCAAACTAGTGGTGTTTCAGGGTGAACACTCAAACGC	988	ималмі с Грнмким
928	TODAAAOTOAAOTAAOATAOOTAAOAOTTOAOAAOAOTA	948	ratmitLDHcloned
			2
	* ** ****** **** ** ** ** * * * ***		
882	TTTAACCCAGAACTGGGCACTGACGACAACGACAGGCAGTGGAAGGAGG	958	MONaemitLDHmRNA
288	TOTGCACACATTTAGGGACTGTAAAATADAAGGAACACTGGAAAGGG		humanmitLDHmRNA
S78	DYSALECCEC SECTION OF THE PROPERTY OF THE PROP		ratmitLDHcloned
760	- ウロ・カー・カー・カー・カー・カー・カー・カー・カー・カー・カー・カー・カー・カー・	クしり	Fower Fruit 14 rm 4 cy

1290 1290 INDADAAGTTDDA-AATAADDDDAAAATDAADATATT		ratmitLDHcloned humanmitLDHMRNA
DTDAADDTATDAADDAADDTAADAATDTADAAA-AATDTA DTDAADDDAADD	8 <i>L</i> 9T	
ATATAOOATTAOATBTBADTATATOABAT-TTTATTOTABBAAAO TTATAAOBTTAOBTBTBATBTTOTBTBTATATATATATTTTATABBADAT	1629	
CTTGTGATATATATATATATATATATATATATATATATAT	1280	
-STASASTEGAS-TEGASSSTTTTTTTTASASSSTASASTASASSTAS	1235	
OTOBETCOTTOAATATATTETCOGTGTGTGTGTGAGCTGCGCTCCCTGCCTCCCTCCCTCCCT	1485	
BTTTADBTDATBTDGTADDGTDADDGTDTDTDTTDTTTTGTADDGTDTDTDTD	7435	
TGCTGGATGTTAATTCTTGTGTGTTCAACTGGTTAGTTGTGAATA TGCTGGATGTTAGTGTGTGTGTGATGTGTGATGTGA	1385	
ADTOCTATATOBACACTAAAATTGTTAAAATCCACACAATAAAATTCCAGACTTAAAATTCTAAAATCCTGACACACAACAACAACAACAACAACAACAACAACAACAAC	1335	

Fig. 13D

mousemitLDHmRNA 1654 TCATGCCAAATAAAACCTTGAACATG

1680

Fig. 14